

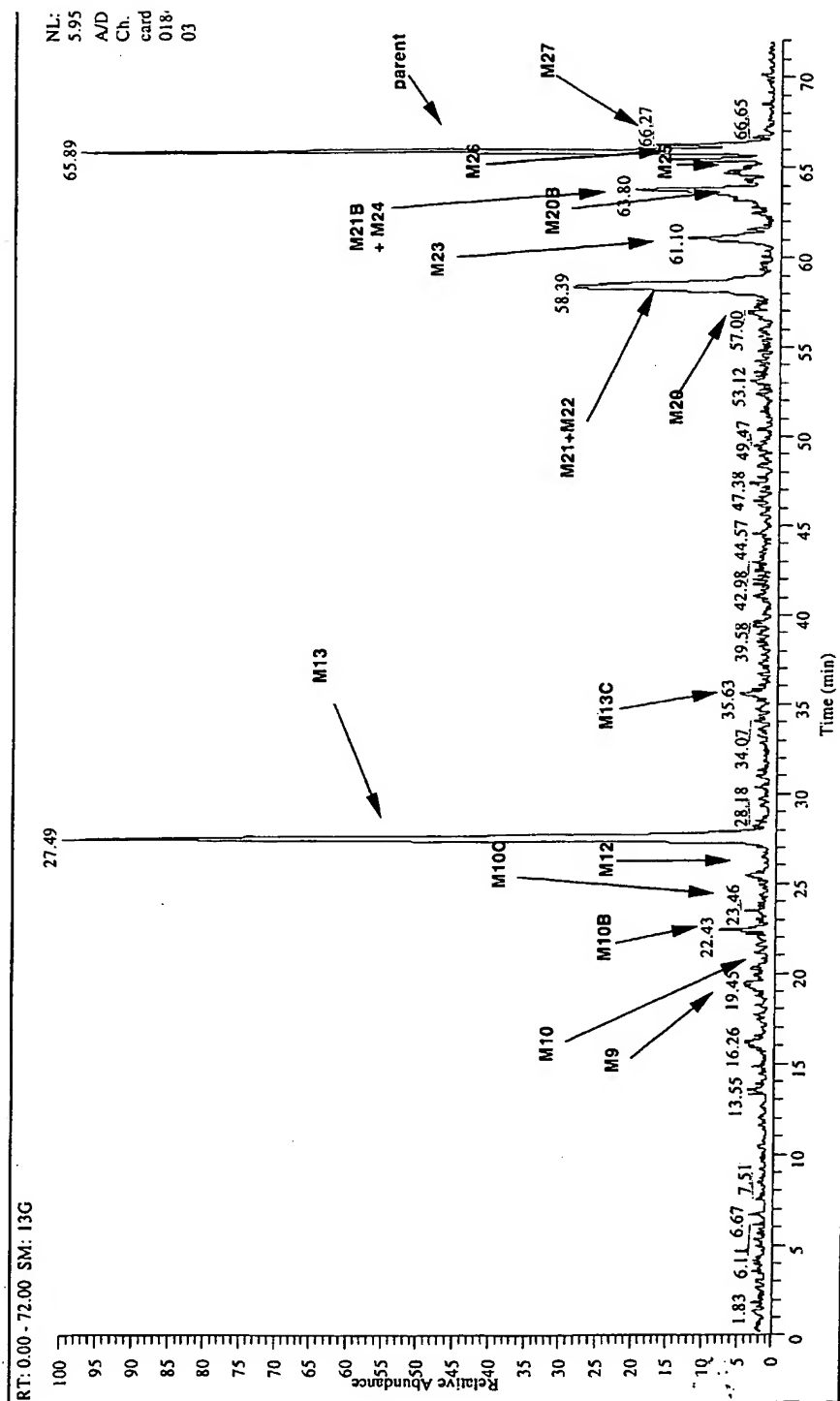
FIGURE 1Radiochromatogram of [¹⁴C]Compound I after 1-hr Incubation with Human Liver Microsomes

Figure 2Radiochromatogram of [14 C]Compound I after 1-hr Incubation with Monkey Liver Microsomes

HA01843\data01843008\01843008.m02

05/24/01 12:33:57 PM

MKLM-TA

RT: 0.00 - 72.00 SM: 13G

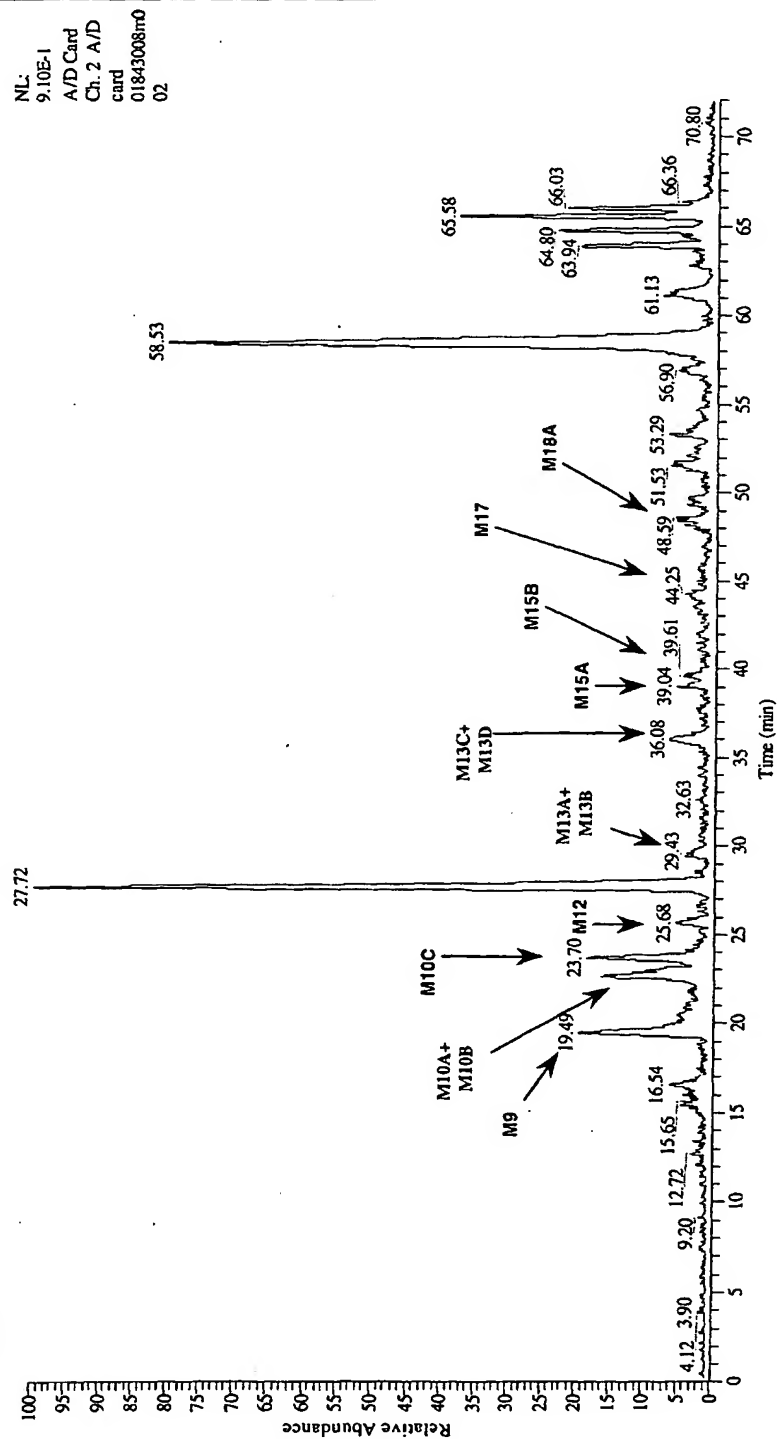


Figure 3

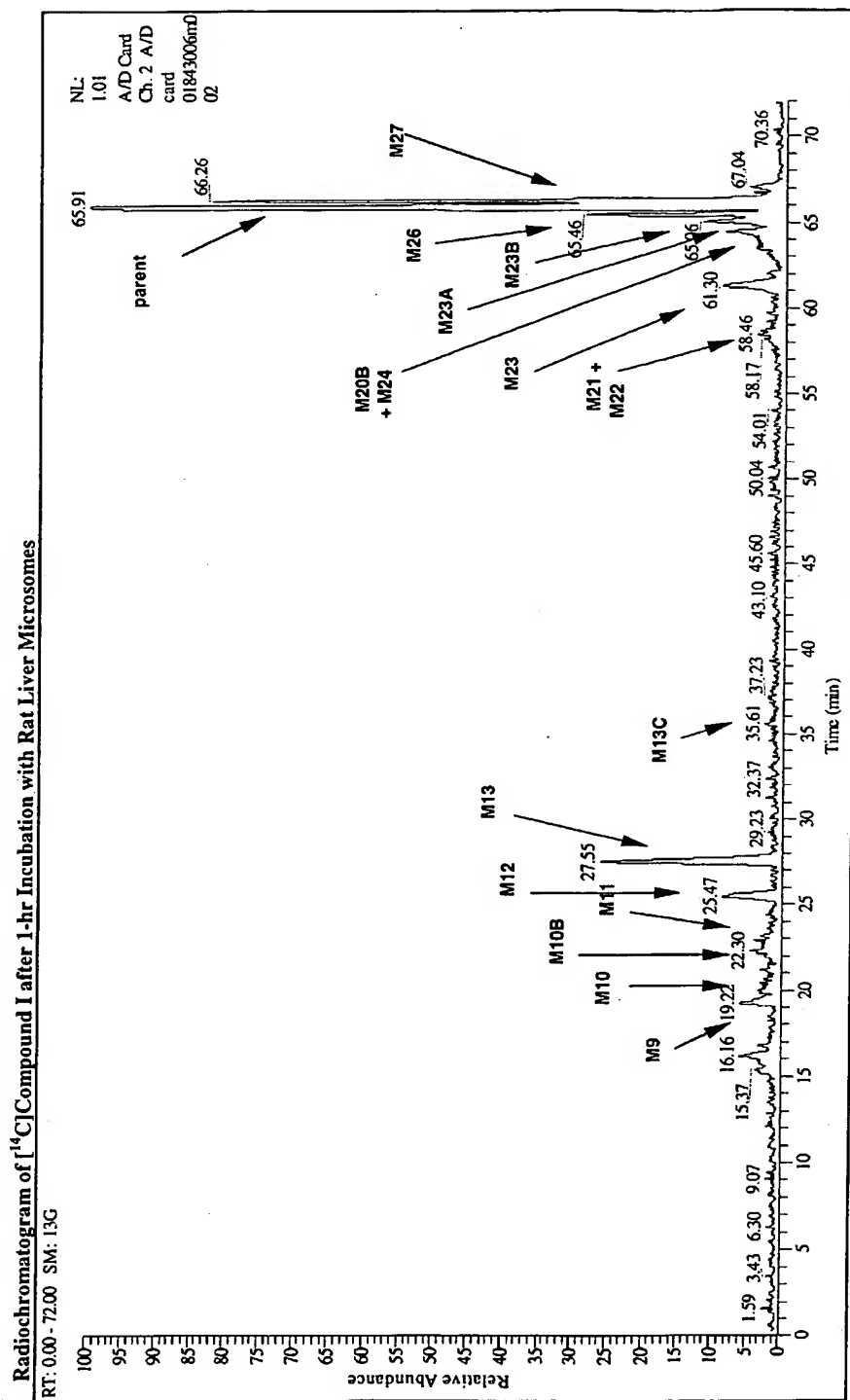
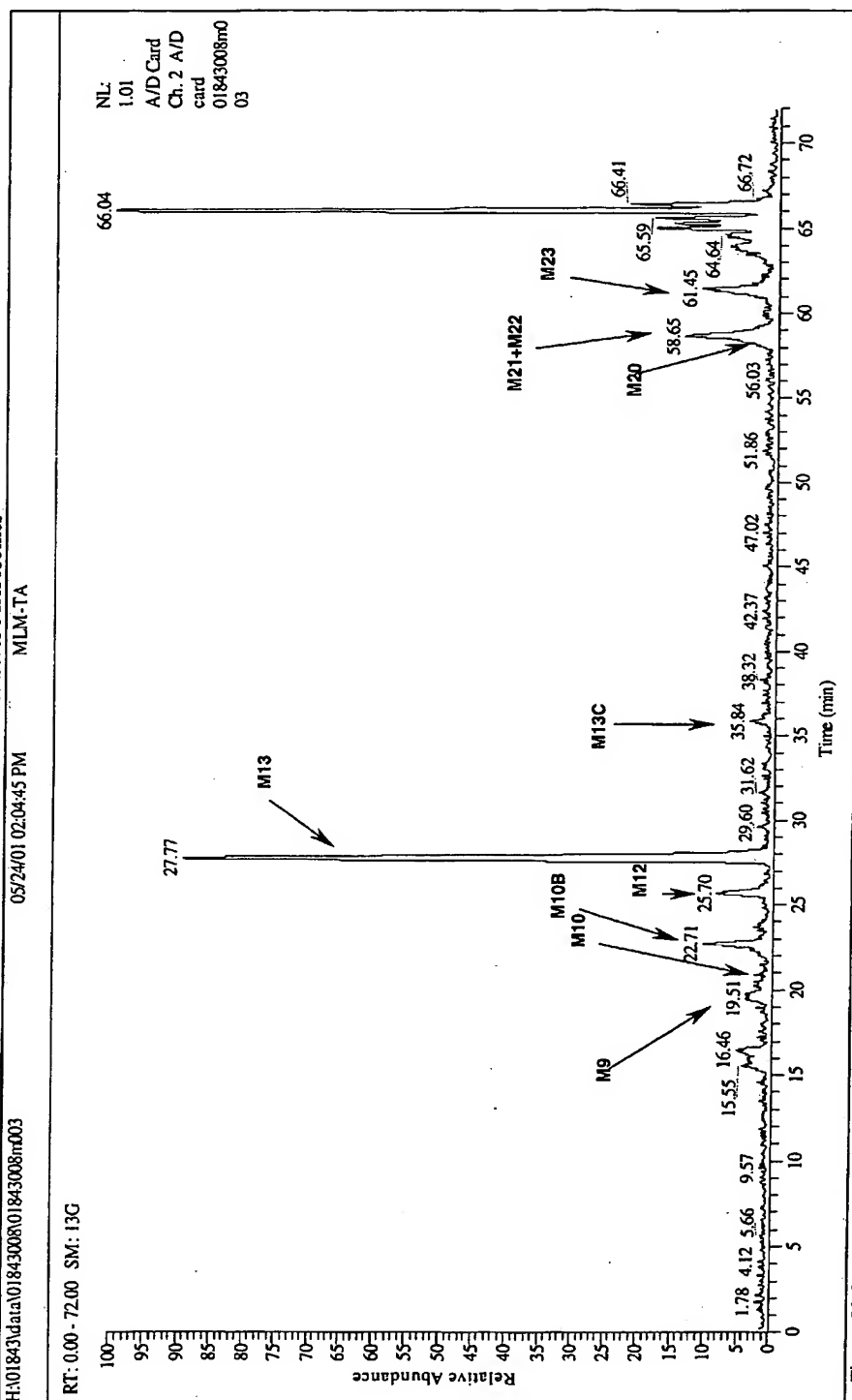


Figure 4
Radiochromatogram of [^{14}C]Compound I after 1-hr Incubation with Mouse Liver Microsomes
H:\0184\data\01843008\01843008m003 MLM-TA
05/24/01 02:04:45 PM



Refer to Figure 50 for the assignment of the metabolite ID of the rest of the peaks.

Figure 5Radiochromatogram of [^{14}C]Compound I after 1-hr Incubation with Dog Liver Microsomes

H:\01843\data\01843008\01843008.m004

05/24/01 03:34:58 PM

DLM-TA

RT: 0.00 - 72.00 SM: 13G

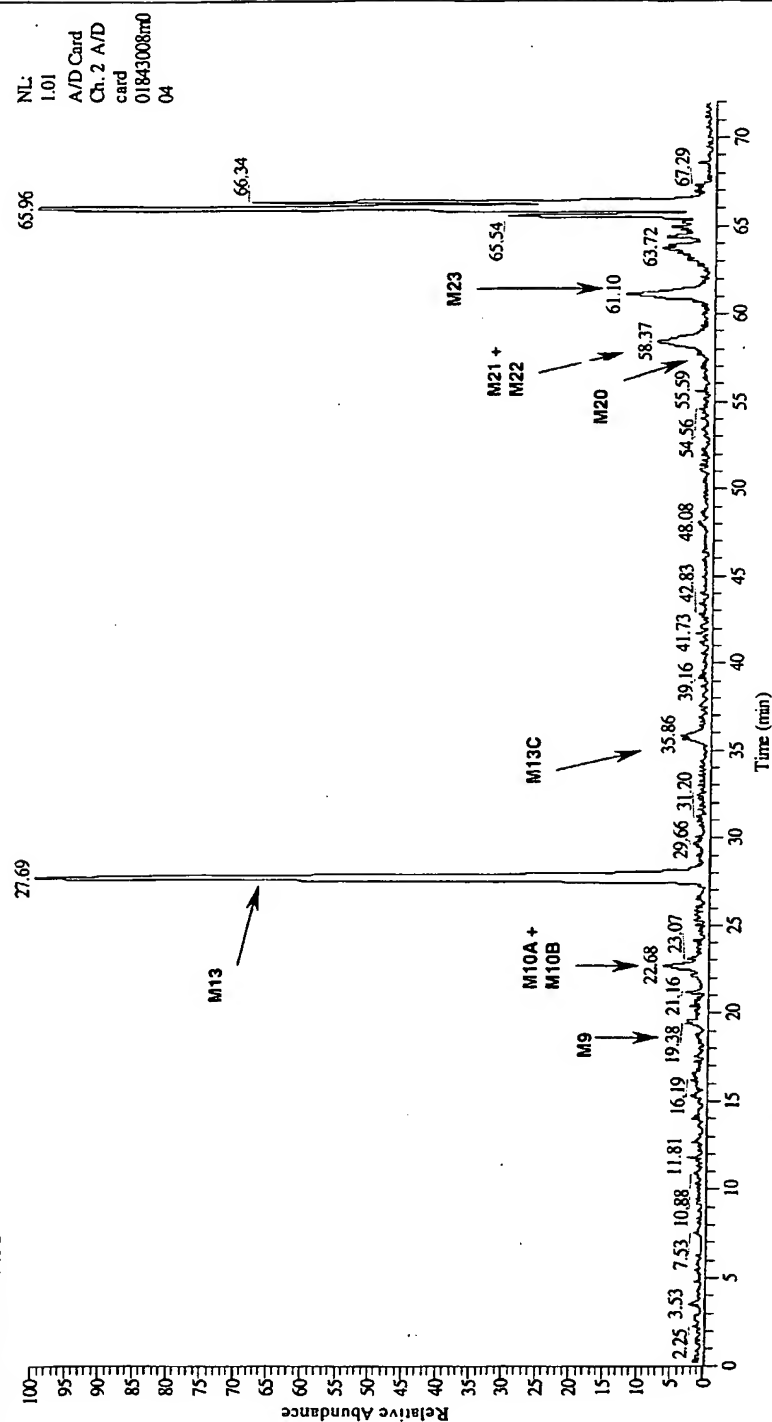
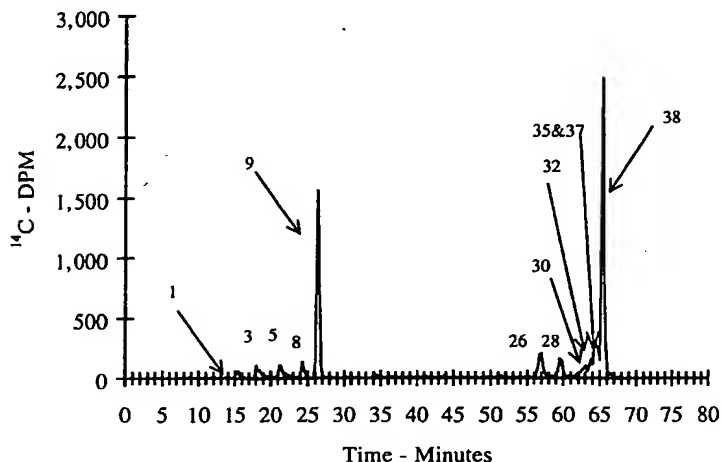


Figure 6

Reconstructed HPLC Radiochromatogram of [^{14}C]Compound I after 1-hr Incubation with Mouse Liver Microsomes



Sample Name: 01843013.D02

Start: 11-May-01

Method: 01843

LSC Reference: LSC977: 101-1

ID: X977:49A (MLM-TA)

Column: Ace 5 C18, 4.6 x 250 mm, 5 μm , #A3538; Guard: RP-8, 4.6 x 30 mm, 5 μm

HPLC System T

Column Temperature: 30°C

Detector: UV at 254 nm; ^{14}C : Raytest

Gradient: 100% A, 3 min to 40% B, 30 min to 60% B (15 min), 10 min to 65% B, 5 min to 100% B (9 min)

A: 0.01 M NH_4OAc (pH=3 w/ HCOOH) B: Methanol Flow Rate: 1.0 mL/min

^{14}C Data Region No.	Metabolite	Start	Stop	R _f	dpm	%ROI	%Total
1		15.25	15.50	15.50	111	1.09%	0.94%
3	M9 + M10	18.00	19.25	18.00	319	3.13%	2.70%
5	M10B	21.00	21.75	21.25	266	2.61%	2.25%
8	M12	24.00	24.50	24.25	229	2.25%	1.94%
9	M13	26.00	27.00	26.25	2706	26.58%	22.91%
26	M20 + M21 + M22	56.25	57.75	57.00	663	6.51%	5.61%
28	M23	59.25	60.25	59.50	451	4.43%	3.82%
30	M20B + M21B + M24	61.75	63.25	63.00	410	4.03%	3.47%
32	M23A	63.50	64.00	63.75	364	3.57%	3.08%
35	M23B+M23C	64.25	64.50	64.50	551	5.41%	4.66%
37	M23D+M26	64.75	65.00	64.75	395	3.88%	3.34%
38	Parent	65.25	66.25	65.25	3717	36.51%	31.47%
DPM Recovered:		11813		DPM in ROI:	10182	100.00%	86.19%
DPM Injected:		12465					
% Recovery:		94.77					

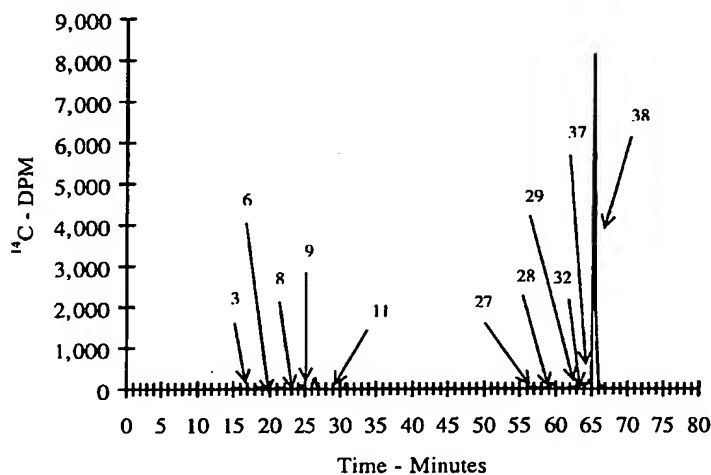
Note: 1) Those peaks which accounted for >0.5% of the total were considered as ROI.

2) M13C and M27 accounted for >0.5% of the total and considered as detected in a trace amount.

7/21

Figure 7

Reconstructed HPLC Radiochromatogram of [¹⁴C]Compound I after 1-hr Incubation with Rat Liver Microsomes



Sample Name: 01843011.D03 Start: 9-May-01
 Method: 01843 LSC Reference: LSC977: 82-83
 ID: X977:48A (RLM-TA)
 Column: Ace 5 C18, 4.6 x 250 mm, 5µm, #A3538; Guard: RP-8, 4.6 x 30 mm, 5µm
 HPLC System T Column Temperature: 30°C
 Detector: UV at 254 nm; ¹⁴C: Raytest
 Gradient: 100% A, 3 min to 40% B, 30 min to 60% B (15 min), 10 min to 65% B, 5 min to 100% B (9 min)
 A: 0.01 M NH₄OAc (pH=3 w/ HCOOH) B: Methanol Flow Rate: 1.0 mL/min

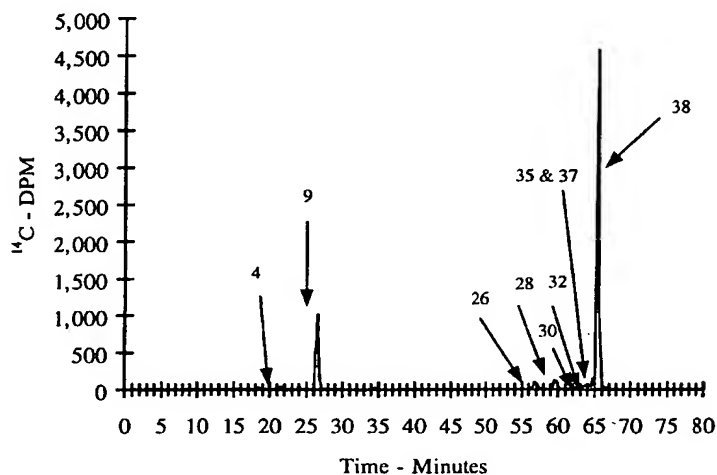
¹⁴ C Data Region No.	Metabolite	Start	Stop	R _t	dpm	%ROI	%Total
3	M9 + M10	18.25	19.25	18.25	183	1.29%	1.16%
6	M10B + M11	21.00	22.00	21.25	151	1.06%	0.96%
8	M12	24.25	24.50	24.25	174	1.23%	1.11%
9	M13	26.25	26.75	26.50	554	3.91%	3.52%
11		30.75	31.00	30.75	173	1.22%	1.10%
27	M21 + M22	56.50	57.25	56.75	159	1.12%	1.01%
28	M23	59.00	60.25	59.50	362	2.55%	2.30%
29	M20B + M24	62.00	63.25	62.75	212	1.49%	1.35%
32	M23A	63.50	64.00	63.75	234	1.65%	1.49%
37	M23B + M26	64.25	65.00	64.75	572	4.03%	3.63%
38	parent	65.25	66.50	65.25	11411	80.44%	72.48%
DPM Recovered:		15744	DPM in ROI:		14185	99.99%	90.11%
DPM Injected:		15460					
% Recovery:		101.84					

Note: 1) Those peaks which accounted for >0.5% of the total were considered as ROI.

2) M13C and M27 accounted for <0.5% of the total and considered as detected in a trace amount.

Figure 8

Reconstructed HPLC Radiochromatogram of [^{14}C]Compound I after 1-hr Incubation with Dog Liver Microsomes



Sample Name: 01843013.D03

Start: 11-May-01

Method: 01843

LSC Reference: LSC977: 114-115

ID: X977:49B (DLM-TA)

Column: Ace 5 C18, 4.6 x 250 mm, 5 μm , #A3538; Guard: RP-8, 4.6 x 30 mm, 5 μm

HPLC System T

Column Temperature: 30°C

Detector: UV at 254 nm; ^{14}C : Raytest

Gradient: 100% A, 3 min to 40% B, 30 min to 60% B (15 min), 10 min to 65% B, 5 min to 100% B (9 min)

A: 0.01 M NH_4OAc (pH=3 w/ HCOOH) B: Methanol Flow Rate: 1.0 mL/min

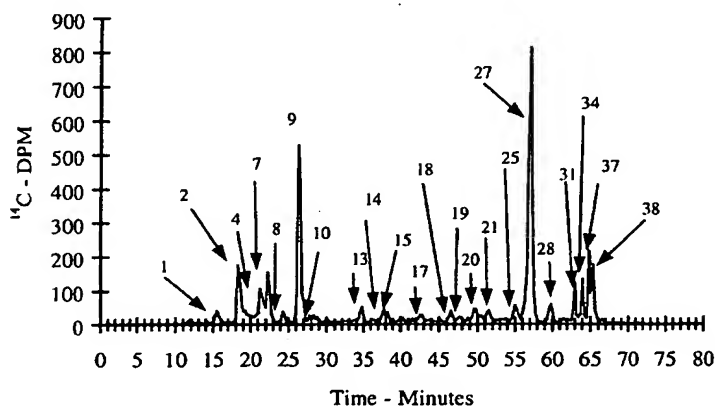
^{14}C Data Region No.	Metabolite	Start	Stop	R_t	dpm	%ROI	%Total
4	M10A + M10B	21.25	21.50	21.50	68	0.72%	0.64%
9	M13	26.25	27.00	26.50	1807	19.15%	17.10%
26	M20 + M21 + M22	56.50	57.25	56.75	255	2.70%	2.41%
28	M23	59.25	60.00	59.50	321	3.40%	3.04%
30	M20B + M21B + M24	62.00	63.25	62.75	256	2.71%	2.42%
32	M23A	63.50	64.00	63.75	195	2.07%	1.85%
35	M23B + M23C + M25	64.25	64.50	64.25	102	1.08%	0.97%
37	M23D + M26	64.75	65.00	64.75	235	2.49%	2.22%
38	parent	65.25	66.50	65.25	6198	65.68%	58.65%
DPM Recovered:		10568	DPM in ROI:		9437	100.00%	89.30%
DPM Injected:		11244					
% Recovery:		93.99					

Note: 1) Those peaks which accounted for >0.5% of the total were considered as ROI.

2) M9, M13C and M27 accounted for <0.5% of the total and considered as detected in a trace amount.

Figure 9

Reconstructed HPLC Radiochromatogram of [^{14}C]Compound I after 1-hr Incubation with Monkey Liver Microsomes



Sample Name: 01843013.D04

Start: 11-May-01

Method: 01843

LSC Reference: LSC977: 105-101

ID: X977:49C (MKLM-TA)

¹⁴ C Data Region No.	Metabolite	Start	Stop	R _t	dpm	%ROI	%Total
1		15.25	15.75	15.50	92	1.37%	1.12%
2	M9	18.00	20.50	18.25	591	8.80%	7.18%
4	M10A + M10B	20.75	22.00	21.25	367	5.46%	4.46%
7	M10C	22.25	22.75	22.25	278	4.14%	3.38%
8	M12	24.25	24.50	24.25	61	0.91%	0.74%
9	M13	26.00	27.00	26.25	1107	16.48%	13.45%
10	M13A + M13B	27.75	28.25	28.25	73	1.09%	0.89%
13	M13C + M13D	34.50	34.75	34.75	88	1.31%	1.07%
14	M15A	37.50	38.00	37.75	95	1.41%	1.15%
15	M15B	38.25	38.50	38.25	56	0.83%	0.68%
17	M17	42.50	42.75	42.75	57	0.85%	0.69%
18	M18A	46.25	46.75	46.50	88	1.31%	1.07%
19	M18B	47.50	47.75	47.75	41	0.61%	0.50%
20	M18C + M19	49.25	50.50	49.75	189	2.81%	2.30%
21	M19A + M19B	51.00	51.75	51.50	119	1.77%	1.45%
25	M20	54.75	55.50	55.00	156	2.32%	1.90%
27	M21 + M22	56.00	57.75	57.00	2176	32.39%	26.44%
28	M23	59.50	60.00	59.75	137	2.04%	1.66%
31	M21B + M24	62.75	63.50	63.00	208	3.10%	2.53%
34	M25	63.75	64.25	64.00	191	2.84%	2.32%
37	M23D + M26	64.75	65.00	64.75	307	4.57%	3.73%
38	Parent	65.25	65.75	65.25	242	3.60%	2.94%
DPM Recovered:		8231	DPM in ROI:		6719	100.01%	81.65%
DPM Injected		8097					
% Recovery:		101.65					

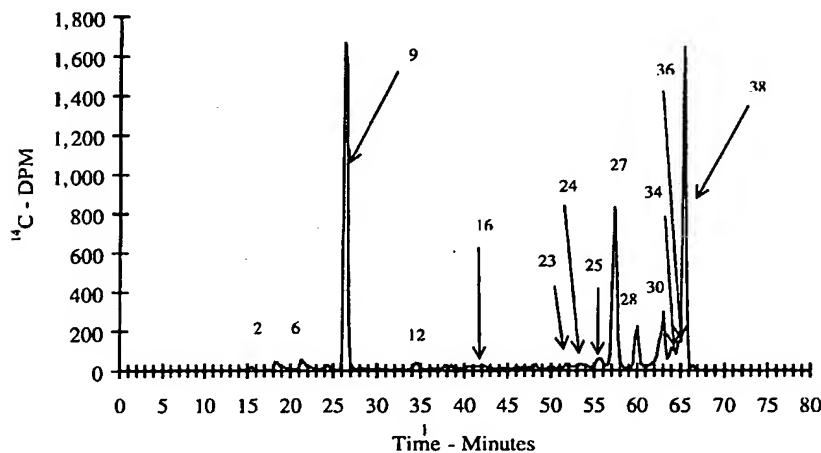
Note: 1) Those peaks which accounted for >0.5% of the total were considered as ROI.

2) M27, and Regions 16, 23, and 24 accounted for <0.5% of the total and considered as detected in a trace amount.

10/21

Figure 10

Reconstructed HPLC Radiochromatogram of [¹⁴C]Compound I after 1-hr Incubation with Human Liver Microsomes



Sample Name: 01843010.D02

Start: 8-May-01

Method: 01843

LSC Reference: LSC977: 88, 90-91

ID: X977:49D (HLM-TA)

Column: Ace 5 C18, 4.6 x 250 mm, 5µm, #A3538; Guard: RP-8, 4.6 x 30 mm, 5µm

HPLC System T

Column Temperature: 30°C

Detector: UV at 254 nm; ¹⁴C: Raytest

Gradient: 100% A, 3 min to 40% B, 30 min to 60% B (15 min), 10 min to 65% B, 5 min to 100% B (15 min)

A: 0.01 M NH₄OAc (pH=3 w/ HCOOH) B: Methanol Flow Rate: 1.0 mL/min

¹⁴ C Data Region No.	Metabolite	Start	Stop	R _f	dpm	%ROI	%Total
2	M9	18.25	18.75	18.25	114	0.97%	0.85%
6	M10B	21.25	21.75	21.25	129	1.10%	0.96%
9	M13	26.00	27.00	26.25	3571	30.53%	26.66%
12	M13C	34.25	34.75	34.50	99	0.85%	0.74%
16		41.25	42.25	42.00	112	0.96%	0.84%
23		51.75	52.25	51.75	88	0.75%	0.66%
24		52.75	54.25	53.50	168	1.44%	1.25%
25	M20	55.00	56.25	55.50	251	2.15%	1.87%
27	M21 + M22	56.50	58.25	57.25	2122	18.14%	15.84%
28	M23	59.50	60.75	60.00	591	5.05%	4.41%
30	M20B + M21B + M24	61.25	63.50	63.00	1044	8.93%	7.79%
34	M25	63.75	64.50	64.00	390	3.33%	2.91%
36	M26	64.75	64.75	64.75	145	1.24%	1.08%
38	parent	65.00	66.00	65.25	2871	24.55%	21.43%
DPM Recovered:		13396	DPM in ROI:		11695	99.99%	87.29%
DPM Injected:		13110					
% Recovery:		102.18					

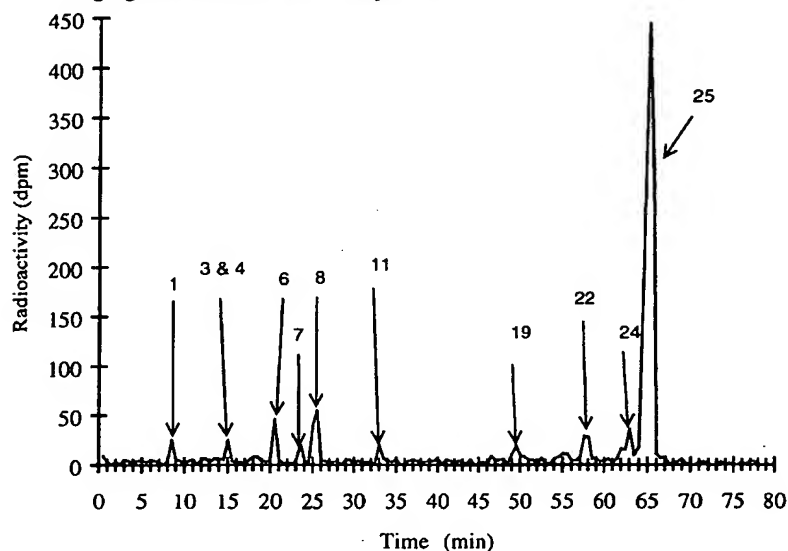
Note: 1) Those peaks which accounted for >0.5% of the total were considered as ROI.

2) Each of M10, M10C, M12, and M27 accounted for <0.5% of the total and considered as detected in trace amounts.

11/21

FIGURE 11

Radiochromatogram of 1 hr Pooled Plasma from Two Rats after a Single 25-mg/kg Oral Dose of [¹⁴C]Compound I



Sample Name: 01842009.D02

Start: 3-Apr-01

Method: 01842

LSC Reference: LSC966:84-87

ID: X966:29E (Pooled Plasma, 1 hr)

Column: Ace 5 C18, 4.6 x 250 mm, 5 μ m, #A3538, Guard: RP-8, 4.6 x 30 mm, 5 μ m

HPLC System T, Column temperature: 30 °C

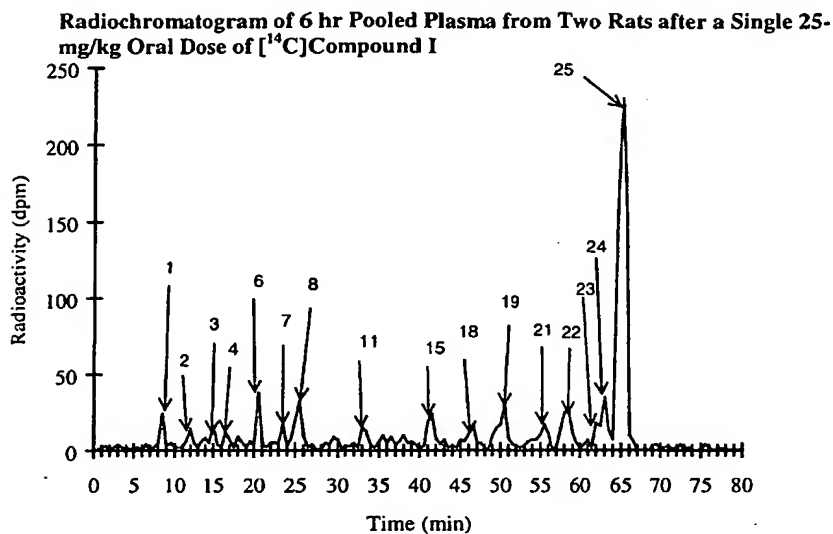
Detector: UV at 254 nm; ¹⁴C: Raytest

Gradient: 100% A, 3 min to 40% B, 30 min to 60% B (15 min), 10 min to 65% B, 5 min to 100% B (15 min)

A: 0.01 M NH₄OAc (pH=3 w/ HCOOH) B: Methanol; Flow Rate: 1.0 mL/min

Region No.	¹⁴ C Data Name	Start	Stop	R _f	dpm	%ROI	%Total
1	M3	8.5	8.5	8.5	25	1.84%	1.35%
3 & 4	M7 + M8 + M8A	15.0	15.0	15.0	25	1.84%	1.35%
6	M10	20.5	20.5	20.5	46	3.39%	2.48%
7	M12	23.5	23.5	23.5	22	1.62%	1.19%
8	M13	25.0	25.5	25.5	96	7.07%	5.18%
11	M14	33.0	33.0	33.0	20	1.47%	1.08%
19		49.5	49.5	49.5	19	1.40%	1.03%
22	M23	57.5	58.0	57.5	57	4.20%	3.08%
24		62.0	63.0	63.0	67	4.94%	3.62%
25	parent	64.0	65.5	65.0	980	72.22%	52.92%
DPM Recovered:		1852		DPM in ROI:	1357	99.99%	73.28%
DPM Injected:		1770		Note: any fraction containing less than 15 dpm was considered as background and not included in the integration.			
% Recovery:		104.63					

12/21

FIGURE 12

Sample Name: 01842009.D03

Start: 9-Apr-01

Method: 01842

LSC Reference: LSC966:89-92

ID: X966:29F (Pooled Plasma, 6 hr)

Region No.	¹⁴ C Data Name	Start	Stop	R _f	dpm	%ROI	%Total
1	M3	8.5	8.5	8.5	24	2.33%	1.57%
2		12.0	12.0	12.0	14	1.36%	0.91%
3	M7	15.0	15.0	15.0	16	1.55%	1.05%
4	M8 + M8A	16.5	16.5	16.5	12	1.16%	0.78%
6	M10	20.5	20.5	20.5	38	3.69%	2.48%
7	M12	23.5	23.5	23.5	17	1.65%	1.11%
8	M13	25.0	25.5	25.5	52	5.04%	3.40%
11	M14	33.0	33.5	33.0	28	2.72%	1.83%
15	M16	41.0	41.5	41.5	43	4.17%	2.81%
18	M19	46.5	46.5	46.5	17	1.65%	1.11%
19		49.5	50.5	50.5	62	6.01%	4.05%
21	M21 or M22	55.5	55.5	55.5	17	1.65%	1.11%
22	M23	57.5	58.5	58.0	64	6.21%	4.18%
23	M24	62.0	62.5	62.0	34	3.30%	2.22%
24		63.0	63.5	63.0	49	4.75%	3.20%
25	parent	64.5	65.5	65.0	544	52.76%	35.53%

DPM Recovered: 1531

DPM in ROI: 1031 100.00% 67.34%

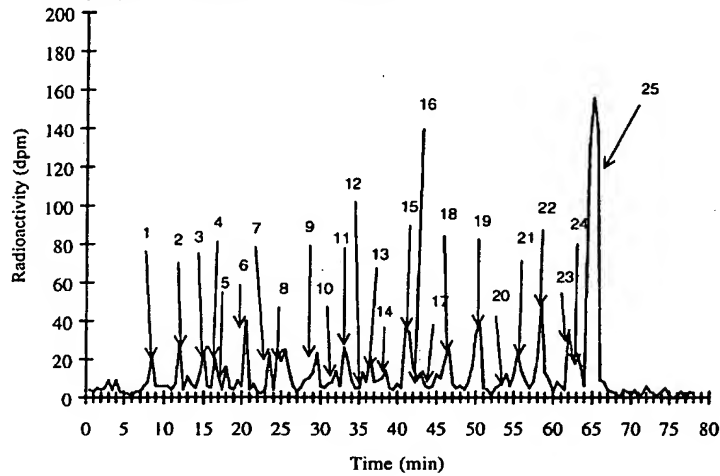
DPM Injected: 1360

% Recovery: 112.57

Note: any fraction containing less than 12 dpm was considered as background and not included in the integration.

FIGURE 13

Radiochromatogram of 12 hr Pooled Plasma from Two Rats after a Single 25mg/kg Oral Dose of [14 C]Compound I



Sample Name: 01842009.D04

Method: 01842

Start: 9-Apr-01

LSC Reference: LSC966:93-96

ID: X966:29G (Pooled Plasma, 12 hr)

14 C Data	Name	Start	Stop	R _f	dpm	%ROI	%Total
1	M3	8.5	8.5	8.5	21	1.59%	1.11%
2		12.0	12.0	12.0	28	2.12%	1.48%
3	M7	14.5	15.0	15.0	35	2.65%	1.85%
4	M8 + M8A	16.5	16.5	16.5	21	1.59%	1.11%
5	M9	18.0	18.0	18.0	16	1.21%	0.84%
6	M10	20.5	20.5	20.5	40	3.03%	2.11%
7	M12	23.5	23.5	23.5	23	1.74%	1.21%
8	M13	24.5	26.0	25.5	83	6.28%	4.38%
9		29.0	29.5	29.5	36	2.73%	1.90%
10		32.0	32.0	32.0	14	1.06%	0.74%
11	M14	33.0	33.5	33.0	45	3.41%	2.37%
12	M15	35.5	35.5	35.5	13	0.98%	0.69%
13		36.5	36.5	36.5	22	1.67%	1.16%
14		38.5	38.5	38.5	14	1.06%	0.74%
15	M16	41.0	42.0	41.0	87	6.59%	4.59%
16		43.0	43.0	43.0	12	0.91%	0.63%
17	M18	45.0	45.0	45.0	12	0.91%	0.63%
18	M19	46.0	46.5	46.5	46	3.48%	2.42%
19		49.5	50.5	50.5	88	6.66%	4.64%
20		54.0	54.0	54.0	12	0.91%	0.63%
21	M21 or M22	55.0	56.0	55.5	50	3.79%	2.64%
22	M23	58.0	59.0	58.5	83	6.28%	4.38%
23	M24	62.0	62.5	62.0	56	4.24%	2.95%
24		63.0	63.5	63.0	40	3.03%	2.11%
25	parent	64.5	65.5	65.0	424	32.10%	22.35%

DPM Recovered: 1897

DPM Injected: 1570

% Recovery: 120.83

DPM in ROI: 1321

100.02%

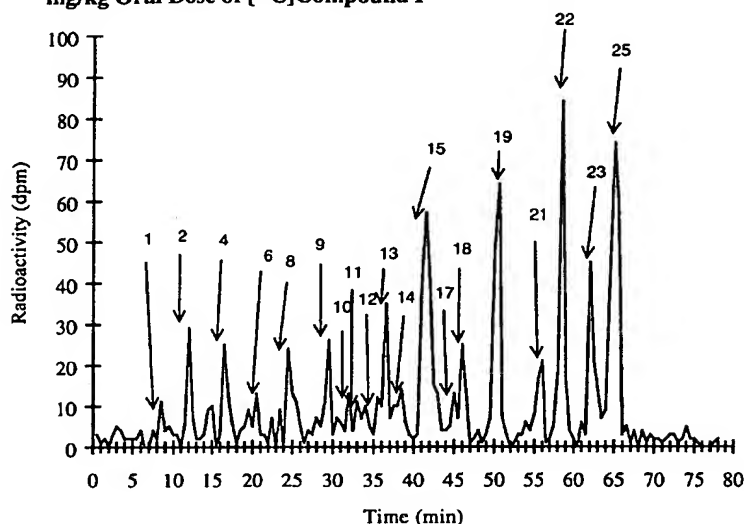
69.66%

Note: any fraction containing less than 12 dpm was considered as background and not included in the integration.

14/21

FIGURE 14

Radiochromatogram of 24 hr Pooled Plasma from Two Rats after a Single 25-mg/kg Oral Dose of [14 C]Compound I



Sample Name: 01842009.D05

Start: 9-Apr-01

Method: 01842

LSC Reference: LSC966:97-101

ID: X966:29H (Pooled Plasma, 24 hr)

Region No.	14 C Data Name	Start	Stop	R _f	dpm	%ROI	%Total
1	M3	8.5	8.5	8.5	11	1.10%	0.77%
2		12.0	12.0	12.0	29	2.89%	2.02%
4	M8 + M8A	16.5	17.0	16.5	37	3.69%	2.58%
6	M10	20.5	20.5	20.5	13	1.30%	0.91%
8	M13	24.5	25.5	24.5	48	4.79%	3.34%
9		29.0	29.5	29.5	37	3.69%	2.58%
10		32.0	32.0	32.0	13	1.30%	0.91%
11	M14	33.0	33.0	33.0	11	1.10%	0.77%
12	M15	35.5	35.5	35.5	12	1.20%	0.84%
13		36.5	36.5	36.5	35	3.49%	2.44%
14		38.5	38.5	38.5	14	1.40%	0.98%
15	M16	41.0	43.0	41.5	166	16.57%	11.57%
17	M18	45.0	45.0	45.0	13	1.30%	0.91%
18	M19	46.0	46.5	46.0	38	3.79%	2.65%
19		50.0	50.5	50.5	113	11.28%	7.87%
21	M21 or M22	55.5	56.0	56.0	38	3.79%	2.65%
22	M23	58.0	59.0	58.5	119	11.88%	8.29%
23	M24	62.0	63.0	62.0	80	7.98%	5.57%
25	parent	64.5	65.5	65.0	175	17.47%	12.20%

DPM Recovered: 1435

DPM in ROI: 1002 100.01%

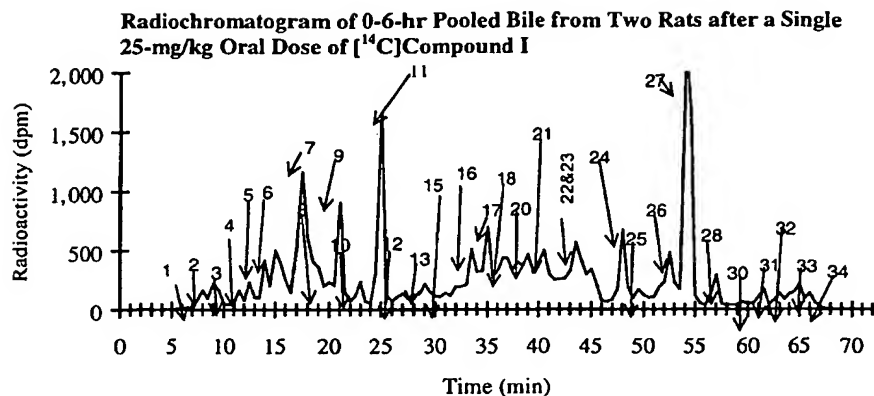
69.85%

DPM Injected: 1220

Note: any fraction containing less than 11 dpm was considered as background and not included in the integration.

% Recovery: 117.62

FIGURE 15



Sample Name: 01842006.D02

Start: 30-Mar-01

Method: 01842

LSC Reference: X966:37-40

ID: Pooled Bile 0-6 hr (X966:17A)

Region No.	¹⁴ C Data Name	Start	Stop	R _f	dpm	%ROI	%Total	% Dose
1	M1 + M2	7.0	8.5	8.0	367	1.16%	1.16%	0.16%
2	M4 + M5	9.0	10.5	9.0	473	1.50%	1.49%	0.20%
3		11.0	12.0	11.5	296	0.94%	0.93%	0.12%
4		12.5	13.5	12.5	436	1.38%	1.37%	0.18%
5	M6	14.0	14.5	14.0	617	1.95%	1.95%	0.26%
6	M7 + M8 + M8A	15.0	16.5	15.0	1303	4.12%	4.11%	0.55%
7	M9	17.0	19.0	17.5	3037	9.60%	9.58%	1.28%
8	M10	19.5	20.5	20.0	641	2.03%	2.02%	0.27%
9	M11	21.0	22.0	21.0	1125	3.56%	3.55%	0.48%
10	M12	22.5	24.0	23.0	475	1.50%	1.50%	0.20%
11	M13	24.5	26.0	25.0	2138	6.76%	6.74%	0.90%
12		26.5	27.5	27.0	334	1.06%	1.05%	0.14%
13		28.0	30.0	29.0	730	2.31%	2.30%	0.31%
15		30.5	31.5	31.0	365	1.15%	1.15%	0.15%
16	M14	32.0	34.0	33.5	1425	4.51%	4.49%	0.60%
17	M15	34.5	35.5	35.0	1282	4.05%	4.04%	0.54%
18		36.0	37.5	36.5	1532	4.84%	4.83%	0.65%
20		38.0	39.5	39.0	1545	4.89%	4.87%	0.65%
21	M16	40.0	41.5	40.5	1447	4.58%	4.56%	0.61%
22 & 23	M17 + M18	42.0	46.5	43.5	2831	8.95%	8.93%	1.20%
24	M19	47.0	49.0	48.0	1224	3.87%	3.86%	0.52%
25		49.5	50.5	49.5	390	1.23%	1.23%	0.16%
26	M20	51.0	53.5	52.5	1412	4.46%	4.45%	0.60%
27	M22	54.0	56.0	54.0	4034	12.76%	12.72%	1.70%
28	M23	56.5	59.0	57.0	587	1.86%	1.85%	0.25%
30		59.5	60.0	59.5	106	0.34%	0.33%	0.04%
31	M24	60.5	62.0	61.5	370	1.17%	1.17%	0.16%
32		62.5	63.5	63.0	302	0.95%	0.95%	0.13%
33	parent	64.0	65.5	65.0	586	1.85%	1.85%	0.25%
34		66.0	67.0	66.0	214	0.68%	0.67%	0.09%

DPM Recovered: 31713

DPM in ROI: 31624

100.01%

99.70%

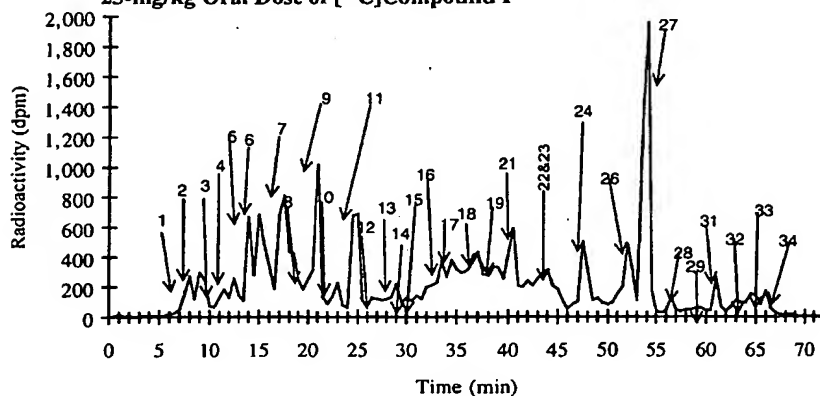
13.35%

DPM Injected: 32752

% Recovery: 96.83

FIGURE 16

Radiochromatogram of 6-12-hr Pooled Bile from Two Rats after a Single 25-mg/kg Oral Dose of [¹⁴C]Compound I



Sample Name: 01842006.D03

Start: 30-Mar-01

Method: 01842

LSC Reference: LSC966:41-44

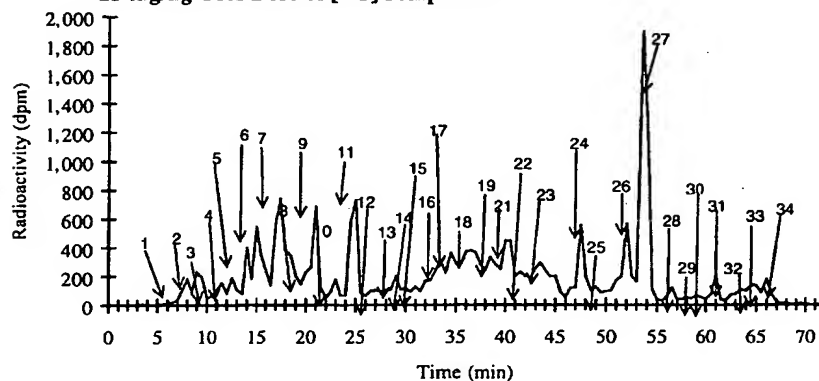
ID: Pooled Bile 6-12 hr (X966:17B)

Region No.	¹⁴ C Data Name	Start	Stop	R _f	dpm	%ROI	%Total	% Dose
1	M1 + M2	6.5	8.5	8.0	612	2.09%	2.08%	0.15%
2	M4 + M5	9.0	10.5	9.0	678	2.32%	2.30%	0.16%
3		11.0	12.0	11.5	442	1.51%	1.50%	0.11%
4		12.5	13.5	12.5	511	1.75%	1.74%	0.12%
5	M6	14.0	14.5	14.0	947	3.24%	3.22%	0.23%
6	M7 + M8 + M8A	15.0	16.5	15.0	1704	5.83%	5.79%	0.41%
7	M9	17.0	19.0	17.5	2596	8.87%	8.82%	0.62%
8	M10	19.5	20.5	20.0	745	2.55%	2.53%	0.18%
9	M11	21.0	22.0	21.0	1231	4.21%	4.18%	0.29%
10	M12	22.5	24.0	23.0	504	1.72%	1.71%	0.12%
11	M13	24.5	26.0	25.0	1546	5.28%	5.25%	0.37%
12		26.5	27.5	26.5	351	1.20%	1.19%	0.08%
13		28.0	29.5	29.0	562	1.92%	1.91%	0.13%
14		30.0	30.5	30.0	233	0.80%	0.79%	0.06%
15		31.0	31.5	31.0	260	0.89%	0.88%	0.06%
16	M14	32.0	34.0	33.5	1282	4.38%	4.36%	0.31%
17	M15	34.5	35.5	34.5	990	3.38%	3.36%	0.24%
18		36.0	37.5	37.0	1392	4.76%	4.73%	0.33%
19		38.0	39.5	38.5	1197	4.09%	4.07%	0.28%
21	M16	40.0	41.5	40.5	1422	4.86%	4.83%	0.34%
22 & 23	M17 + M18	42.0	46.0	44.0	1858	6.35%	6.31%	0.44%
24	M19	46.5	50.0	47.5	1403	4.80%	4.77%	0.33%
26	M20	50.5	53.0	52.0	1372	4.69%	4.66%	0.33%
27	M22	53.5	55.5	54.0	3479	11.89%	11.82%	0.83%
28	M23	56.0	57.5	56.5	260	0.89%	0.88%	0.06%
29		58.0	60.0	59.0	259	0.89%	0.88%	0.06%
31	M24	60.5	62.0	61.0	412	1.41%	1.40%	0.10%
32		62.5	63.5	63.0	273	0.93%	0.93%	0.07%
33	parent	64.0	65.5	64.5	475	1.62%	1.61%	0.11%
34		66.0	67.0	66.0	257	0.88%	0.87%	0.06%
DPM Recovered:		29429	DPM in ROI:		29253	100.00%	99.37%	6.98%
DPM Injected:		30378						
% Recovery:		96.88						

17/21

FIGURE 17

Radiochromatogram of 12-24-hr Pooled Bile from Two Rats after a Single 25-mg/kg Oral Dose of [¹⁴C]Compound I



Sample Name: 01842006.D04

Method: 01842

ID: Pooled Bile 12-24 hr (X966:17C)

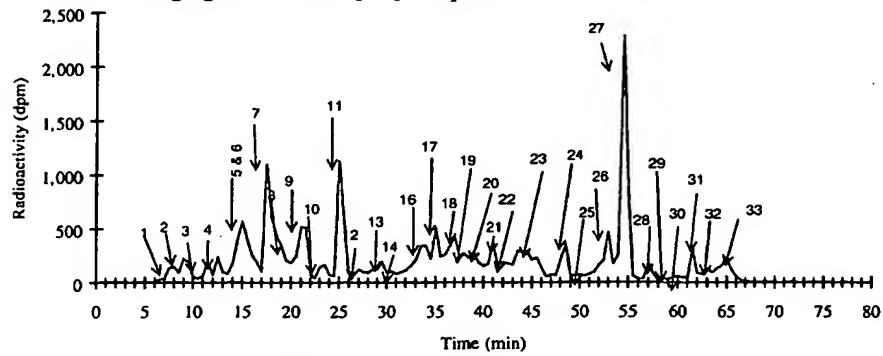
Start: 30-Mar-01

LSC Reference: LSC966:45-48

Region No.	¹⁴ C Data Name	Start	Stop	R _f	dpm	%ROI	%Total	% Dose
1	M1 + M2	6.5	8.5	8.0	434	1.68%	1.68%	0.09%
2	M4 + M5	9.0	10.0	9.0	467	1.81%	1.80%	0.10%
3		10.5	12.0	11.5	394	1.53%	1.52%	0.08%
4		12.5	13.5	12.5	375	1.45%	1.45%	0.08%
5	M6	14.0	14.5	14.0	581	2.25%	2.24%	0.12%
6	M7 + M8 + M8A	15.0	16.5	15.0	1273	4.94%	4.92%	0.27%
7	M9	17.0	19.0	17.5	2229	8.65%	8.61%	0.46%
8	M10	19.5	20.5	20.5	633	2.46%	2.45%	0.13%
9	M11	21.0	22.0	21.0	856	3.32%	3.31%	0.18%
10	M12	22.5	24.0	23.0	404	1.57%	1.56%	0.08%
11	M13	24.5	26.0	25.0	1454	5.64%	5.62%	0.30%
12		26.5	27.5	27.0	297	1.15%	1.15%	0.06%
13		28.0	29.5	29.0	510	1.98%	1.97%	0.11%
14		30.0	30.5	30.0	207	0.80%	0.80%	0.04%
15		31.0	31.5	31.0	222	0.86%	0.86%	0.05%
16	M14	32.0	34.0	33.5	1107	4.30%	4.28%	0.23%
17	M15	34.5	35.5	34.5	942	3.65%	3.64%	0.20%
18		36.0	38.0	36.5	1642	6.37%	6.34%	0.34%
19		38.5	39.5	38.5	864	3.35%	3.34%	0.18%
21	M16	40.0	41.0	40.0	1093	4.24%	4.22%	0.23%
22	M17	41.5	42.5	41.5	618	2.40%	2.39%	0.13%
23	M18	43.0	46.0	43.5	1328	5.15%	5.13%	0.28%
24	M19	46.5	48.5	47.5	1087	4.22%	4.20%	0.23%
25		49.0	49.5	49.0	208	0.81%	0.80%	0.04%
26	M20	50.0	53.0	52.0	1483	5.75%	5.73%	0.31%
27	M22	53.5	55.5	53.5	3244	12.59%	12.53%	0.68%
28	M23	56.0	57.5	56.5	255	0.99%	0.99%	0.05%
29		58.0	58.5	58.0	93	0.36%	0.36%	0.02%
30		59.0	60.0	59.0	156	0.61%	0.60%	0.03%
31	M24	60.5	62.0	61.0	338	1.31%	1.31%	0.07%
32		62.5	64.0	63.5	348	1.35%	1.34%	0.07%
33	parent	64.5	65.5	64.5	359	1.39%	1.39%	0.08%
34		66.0	67.0	66.0	273	1.06%	1.05%	0.06%
DPM Recovered:		25882	DPM in ROI:		25774	99.99%	99.58%	5.38%
DPM Injected:		26360						
% Recovery:		98.19						

FIGURE 18

Radiochromatogram of 0-24-hr Pooled Bile from Two Rats after a Single 25-mg/kg Oral Dose of [¹⁴C]Compound I



Sample Name: 01842008.D03

Method: 01842

ID: X966:28B (Pooled Bile, 0-24 hr)

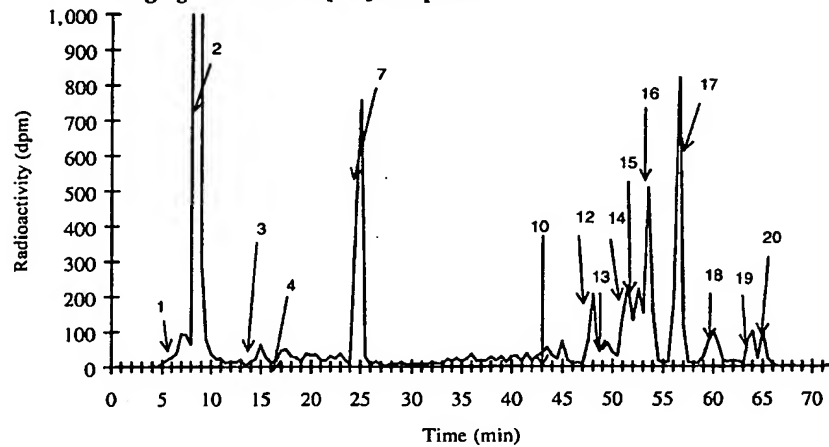
Start: 3-Apr-01

LSC Reference: LSC966:80-83

Region No.	¹⁴ C Data Name	Start	Stop	R _f	dpm	%ROI	%Total	% Dose
1	M1 + M2	6.5	8.5	8.0	442	1.68%	1.68%	0.43%
2	M4 + M5	9.0	10.5	9.0	502	1.91%	1.90%	0.49%
3		11.0	12.0	11.5	329	1.25%	1.25%	0.32%
4		12.5	13.5	12.5	414	1.58%	1.57%	0.41%
5 & 6	M6 + M7 + M8 + M8A	14.0	17.0	15.0	2111	8.04%	8.00%	2.06%
7	M9	17.5	19.5	17.5	2671	10.17%	10.12%	2.61%
8	M10	20.0	20.5	20.5	432	1.64%	1.64%	0.42%
9	M11	21.0	22.5	21.0	1141	4.34%	4.32%	1.11%
10	M12	23.0	24.5	23.5	439	1.67%	1.66%	0.43%
11	M13	25.0	26.5	25.0	1853	7.05%	7.02%	1.81%
12		27.0	28.0	27.0	312	1.19%	1.18%	0.30%
13		28.5	30.0	29.5	536	2.04%	2.03%	0.52%
14		30.5	31.0	30.5	182	0.69%	0.69%	0.18%
16	M14	31.5	34.5	34.0	1477	5.62%	5.60%	1.44%
17	M15	35.0	35.5	35.0	748	2.85%	2.84%	0.73%
18		36.0	37.5	37.0	1269	4.83%	4.81%	1.24%
19		38.0	38.5	38.0	501	1.91%	1.90%	0.49%
20		39.0	40.0	39.0	612	2.33%	2.32%	0.60%
21	M16	40.5	41.5	41.0	693	2.64%	2.63%	0.68%
22	M17	42.0	43.0	42.0	537	2.04%	2.04%	0.53%
23	M18	43.5	46.5	43.5	1500	5.71%	5.69%	1.47%
24	M19	47.0	49.0	48.5	757	2.88%	2.87%	0.74%
25		49.5	50.5	50.0	198	0.75%	0.75%	0.19%
26	M20	51.0	53.5	53.0	1174	4.47%	4.45%	1.15%
27	M22	54.0	56.0	54.5	3444	13.11%	13.05%	3.37%
28	M23	56.5	58.0	57.0	307	1.17%	1.16%	0.30%
29		58.5	59.0	58.5	61	0.23%	0.23%	0.06%
30		59.5	60.5	60.0	157	0.60%	0.60%	0.15%
31	M24	61.0	62.5	61.5	527	2.01%	2.00%	0.52%
32		63.0	63.5	63.0	223	0.85%	0.85%	0.22%
33	parent	64.0	66.5	65.0	720	2.74%	2.73%	0.70%
DPM Recovered:		26382	DPM in ROI:		26269	99.99%	99.58%	25.67%
DPM Injected:		24582						
% Recovery:		107.32						

FIGURE 19

Radiochromatogram of 0-12-hr Pooled Urine from Two Rats after a Single 25-mg/kg Oral Dose of [^{14}C]Compound I



Sample Name: 01842006.D05

Start: 30-Mar-01

Method: 01842

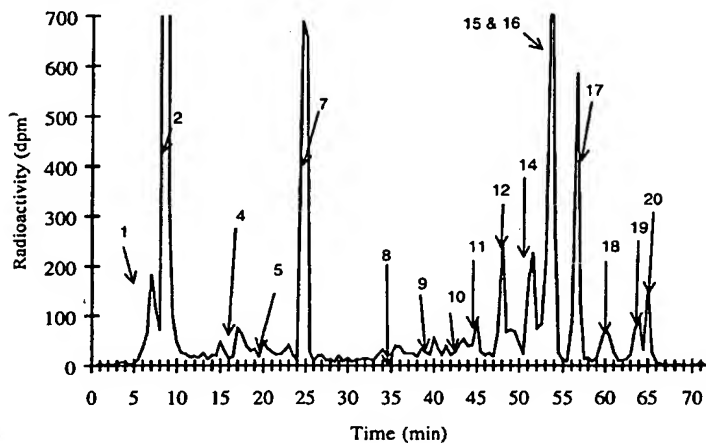
LSC Reference: LSC966:49-52

ID: Pooled Urine 0-12hr (X966:17D)

Region No.	^{14}C Data Name	Start	Stop	R_f	dpm	%ROI	%Total	% Dose
1		6.0	8.0	7.0	310	2.24%	2.05%	0.06%
2	M3	8.5	10.0	8.5	8223	59.55%	54.41%	1.47%
3	M7 + M8 + M8A	15.0	15.5	15.0	90	0.65%	0.60%	0.02%
4	M9	17.0	18.5	17.5	151	1.09%	1.00%	0.03%
7	M13	24.5	25.5	25.0	1181	8.55%	7.81%	0.21%
10	M17	42.5	44.0	43.5	151	1.09%	1.00%	0.03%
12	M19	47.5	49.0	48.0	381	2.76%	2.52%	0.07%
13		49.5	50.5	49.5	135	0.98%	0.89%	0.02%
14		51.0	52.0	51.5	499	3.61%	3.30%	0.09%
15	M20	52.5	53.0	52.5	371	2.69%	2.45%	0.07%
16	M21	53.5	54.0	53.5	660	4.78%	4.37%	0.12%
17	M23	56.0	57.0	56.5	1113	8.06%	7.36%	0.20%
18	M24	59.0	60.5	60.0	267	1.93%	1.77%	0.05%
19		63.5	64.0	64.0	174	1.26%	1.15%	0.03%
20	parent	65.0	65.0	65.0	103	0.75%	0.68%	0.02%
DPM Recovered:		15114	DPM in ROI:		13809	99.99%	91.36%	2.49%
DPM Injected:		16580						
% Recovery:		91.16						

FIGURE 20

Radiochromatogram of 12-24-hr Pooled Urine from Two Rats after a Single 25-mg/kg Oral Dose of [¹⁴C]Compound I



Sample Name: 01842006.D06

Start: 30-Mar-01

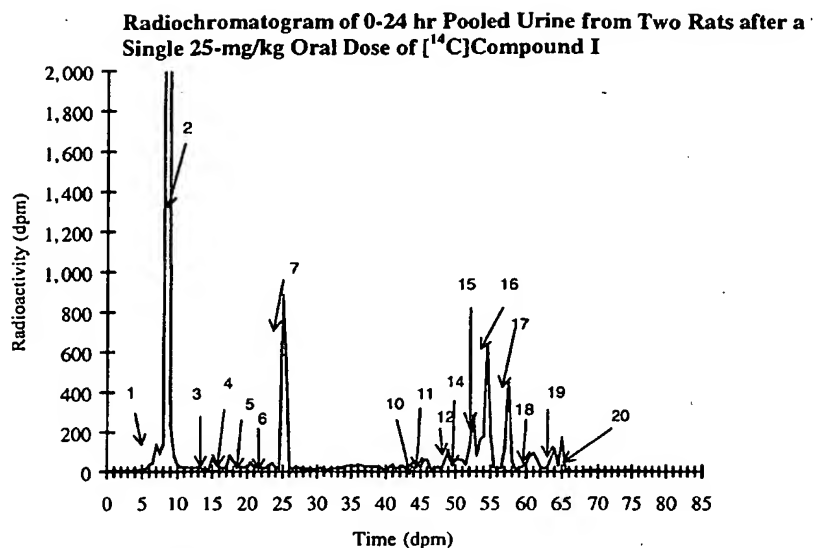
Method: 01842

LSC Reference: LSC966:53-56

ID: Pooled Urine 12-24hr (X966:17E)

Region No.	¹⁴ C Data Name	Start	Stop	R _f	dpm	%ROI	%Total	% Dose
1		6.0	8.0	7.0	466	3.04%	2.82%	0.03%
2	M3	8.5	10.5	8.5	8781	57.31%	53.09%	0.64%
4	M9	17.0	19.0	17.0	242	1.58%	1.46%	0.02%
5	M10	20.0	21.0	20.0	108	0.70%	0.65%	0.01%
7	M13	24.5	25.5	24.5	1370	8.94%	8.28%	0.10%
8	M15	35.5	37.0	35.5	127	0.83%	0.77%	0.01%
9	M16	40.0	40.5	40.0	91	0.59%	0.55%	0.01%
10	M17	42.5	44.0	43.5	161	1.05%	0.97%	0.01%
11	M18	44.5	45.5	45.0	157	1.02%	0.95%	0.01%
12	M19	47.5	50.0	48.0	558	3.64%	3.37%	0.04%
14		51.0	52.0	51.5	456	2.98%	2.76%	0.03%
15 & 16	M20 + M21	52.5	54.0	53.5	1431	9.34%	8.65%	0.10%
17	M23	56.0	57.0	56.5	794	5.18%	4.80%	0.06%
18	M24	59.5	61.0	60.0	204	1.33%	1.23%	0.01%
19		63.5	64.5	64.0	196	1.28%	1.19%	0.01%
20	parent	65.0	65.5	65.0	181	1.18%	1.09%	0.01%
DPM Recovered:		16540	DPM in ROI:		15323	99.99%	92.63%	1.10%
DPM Injected:		16975						
% Recovery:		97.44						

FIGURE 21



Sample Name: 01842008.D02

Start: 3-Apr-01

Method: 01842

LSC Reference: LSC966:76-79

ID: X966:28A (Pooled Urine, 0-24 hr)

Region No.	14 C Data Name	Start	Stop	R _f	dpm	%ROI	%Total	%Dose
1		6.0	7.5	7.0	306	2.02%	1.91%	0.07%
2	M3	8.0	13.0	8.5	8998	59.49%	56.18%	2.19%
3	M7 + M8 + M8A	15.0	16.0	15.0	120	0.79%	0.75%	0.03%
4	M9	17.0	19.5	17.5	238	1.57%	1.49%	0.06%
5	M10	20.0	22.5	20.5	145	0.96%	0.91%	0.04%
6	M12	23.0	24.0	23.5	93	0.61%	0.58%	0.02%
7	M13	25.0	26.0	25.0	1409	9.32%	8.80%	0.34%
10	M17	42.0	45.0	44.0	229	1.51%	1.43%	0.06%
11	M18	45.5	46.0	46.0	118	0.78%	0.74%	0.03%
12	M19	48.5	49.5	49.0	201	1.33%	1.26%	0.05%
14		50.0	51.5	50.5	209	1.38%	1.31%	0.05%
15	M20	52.0	53.0	52.5	476	3.15%	2.97%	0.12%
16	M21	53.5	56.0	54.5	1197	7.91%	7.47%	0.29%
17	M23	56.5	58.0	57.5	629	4.16%	3.93%	0.15%
18	M24	59.0	62.5	61.0	356	2.35%	2.22%	0.09%
19		63.0	64.5	64.0	231	1.53%	1.44%	0.06%
20	parent	65.0	65.0	65.0	169	1.12%	1.06%	0.04%
DPM Recovered:		16015	DPM in ROI:		15124	99.98%	94.45%	3.69%
DPM Injected:		17175						
% Recovery:		93.25						